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wherein Z is C=O or a covalent band; X is H or  $O(C_1-C_4)$  alkyl,  $R^1$  and  $R^2$  are individually  $(C_1-C_4)$  alkyl or together with N are a saturated heterocyclic group,  $R^3$  is ethyl or chloroethyl,  $R^4$  is H or together with  $R^3$  is  $-CH_2-CH_2$ - or -S-,  $R^5$  is I,  $O(C_1-C_4)$  alkyl or H and  $R^6$  is I,  $O(C_1-C_4)$  alkyl or H; or a pharmaceutically acceptable salt thereof.

- 159. (New) The method of claim 158 wherein the compound of formula (I) is tamoxifen or a pharmaceutically acceptable salt thereof.
- 160. (New) The method of claim 158 wherein the compound of formula (I) is idoxifene or a pharmaceutically acceptable salt thereof.
- 161. (New) The method of claim 158 wherein the compound of formula (I) is toremifene or a pharmaceutically acceptable salt thereof.
- 162. (New) The method of claim 158 wherein the administration is to a human patient.
- 163. (New) The method of claim 158 wherein the administration is before, during or after said procedure.

- 164. (New) The method of claim 158 wherein the administration is in/a series of spaced doses.
- 165. (New) The method of claim 158 wherein the administration/is parenteral.
- 166. (New) The method of claim 158 wherein the administration is oral.
- 167. (New) The method of claim 158 wherein the administration is systemic.
- 168. (New) The method of claim 158 wherein the compound of formula (I) is administered via a sustained release dosage form.
- 169. (New) The method of claim 158 wherein the administration is localized at the site of the vascular trauma.
- 170. (New) The method of claim 158 wherein the compound directly or indirectly increases the level of active TGF-beta.
- 171. (New) The method of claim 158 wherein the compound of formula (I) is raloxifene, or a pharmaceutically acceptable salt thereof.
- 172. (New) The method of claim 1/8 wherein the compound of formula (I) is droloxifene, or a pharmaceutically acceptable salt thereof.
- 173. (New) A therapeutic method for preventing or treating a cardiovascular or vascular indication characterized by a decreased lumen diameter comprising administering to a mammal at risk of or afflicted with said cardiovascular or vascular indication, a cytostatic dose of a therapeutic agent, wherein the therapeutic agent is a compound of formula (I):

$$(R^1)(R^2)N(CH_2)_2O$$
 $(Z)$ 
 $R^3$ 
 $(I)$ 

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wherein Z is C=O or a covalent bond; Y is H or  $O(C_1-C_4)$ alkyl,  $R^1$  and  $R^2$  are individually  $(C_1-C_4)$ alkyl or together with N are a saturated heterocyclic group,  $R^3$  is ethyl or chloroethyl,  $R^4$  is H,  $R^5$  is I,  $O(C_1-C_4)$ alkyl or H and  $R^6$  is I,  $O(C_1-C_4)$ alkyl or H with the proviso that when  $R^4$ ,  $R^5$ , and  $R^6$  are H,  $R^3$  is not ethyl; or a pharmaceutically acceptable salt thereof.

- 174. (New) The method of claim 173 wherein the cytostatic dose is effective to increase the level of TGF-beta so as to decrease lesion formation or development, inhibit lipid accumulation, increase plaque stability, maintain or increase vessel lumen diameter, or any combination thereof.
- 175. (New) The method of claim 173 wherein the compound of formula (I) is idoxifene, 4-iodotamoxifen, 3-iodotamoxifen, toremifene, or a pharmaceutically acceptable salt thereof.
- 176. (New) The method of claim 173 wherein the compound of formula (I) is idoxifene or a pharmaceutically acceptable salt thereof.
- 177. (New) The method of claim 173 wherein the compound of formula (I) is toremifene or a pharmaceutically acceptable salt thereof.

- 178. (New) The method of claim 173 wherein the administration is systemic.
- 179. (New) The method of claim 173 wherein the compound of formula (I) is administered via a sustained release dosage form.



- 180. (New) The method of claim 173 wherein the administration is localized at the site of the vascular trauma.
- 181. (New) The method of claim 173 wherein the compound directly or indirectly increases the level of active TGF-beta.
- 182. (New) A therapeutic method of increasing the level of TGF-beta in a mammal in need thereof, comprising administering an effective amount of a compound of formula (I):

$$(R^1)(R^2)N(CH_2)_2O$$
 $(Z)$ 
 $R^3$ 
 $(I)$ 

wherein Z is C=O or a covalent bond; Y is H or  $O(C_1-C_4)$ alkyl,  $R^1$  and  $R^2$  are individually  $(C_1-C_4)$ alkyl or together with N are a saturated heterocyclic group,  $R^3$  is ethyl or chloroethyl,  $R^4$  is H or together with  $R^3$  is  $-CH_2-CH_2$ - or -S-,  $R^5$  is I, OH,  $O(C_1-C_4)$ alkyl or H and  $R^6$  is I,  $O(C_1-C_4)$ alkyl or H with the proviso that when  $R^4$ ,  $R^5$ , and  $R^6$  are H,  $R^3$  is not ethyl; or a pharmaceutically acceptable salt thereof.

- 183. (New) The method of claim 182 wherein the increase in TGF-beta reduces or inhibits diabetic retinopathy.
- 184. (New) The method of claim 182 wherein the mammal is diabetic.
- 185. (New) The method of claim 184 wherein the diabetic has retinopathy.
- 186. (New) The method of claim 182 wherein the compound indirectly or directly increases the level of active TGF-beta in vascular tissue.
- 187. (New) The method of claim 182 wherein the compound is a TGF-beta production stimulator.
- 188. (New) The method of claim 182 wherein the compound is a TGF-beta activator.
- 189. (New) The method of claim 182 wherein the compound increases the production of TGF-beta mRNA.
- 190. (New) The method of claim 182 wherein the compound increases the cleavage of the latent form of TGF-beta.
- 191. (New) The method of claim 182 wherein the compound increases the bioavailability of TGF-beta.
- 192. (New) The method of claim 182 wherein the compound is idoxifene or a pharmaceutically acceptable salt thereof.
- 193. (New) The method of claim 182 wherein the compound is toremifene or a pharmaceutically acceptable salt thereof.

- 194. (New) The method of claim 182 wherein the compound is droloxifene or a pharmaceutically acceptable salt thereof.
- 195. (New) The method of claim 182 wherein the compound is tamoxifen or a pharmaceutically acceptable salt thereof.
- 196. (New) The method of claim 158, 173 or 182 wherein the compound forms cellular DNA adducts at level which is reduced relative to DNA adduct formation by tamoxifen.
- 197. (New) The method of claim 158/173 or 182 wherein the compound has estrogenic activity which is reduced relative to the estrogenic activity of tamoxifen.
- 198. (New) The method of claim 158, 173 or 182 wherein the compound does not form cellular DNA adducts.
- 199. (New) The method of claim 158, 173 or 182 wherein the compound has no estrogenic activity.
- 200. (New) A method of increasing the level of TGF-beta in a mammal in need thereof, comprising administering an effective amount of an agent that directly or indirectly elevates the level of active TGF-beta in said mammal, wherein the agent has reduced estrogenic activity relative to tamoxifen, reduced DNA adduct formation relative to tamoxifen, or any combination thereof.
- 201. (New) The method of claim 200 wherein the agent is a structural analog of tamoxifen or a pharmaceutically acceptable salt thereof.

- 202. (New) The method of claim 200 wherein the agent is idoxifene or a pharmaceutically acceptable salt thereof.
- 203. (New) The method of claim 200 wherein the agent is toremifene or a pharmaceutically acceptable salt thereof.

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- 204. (New) The method of claim 158 wherein the non-aortal smooth muscle cells which are inhibited are present in a non-coronary artery.
  - (New) The method of claim 188, 173, 182, or 200 wherein the administration increases the level of latent TGF-beta relative to the level of latent TGF-beta prior to said administration.
- 206. (New) The method of claim 158, 173, 182, or 200 wherein the administration increases the level of active TGF-beta relative to the level of active TGF-beta prior to said administration.
- 207. (New) A therapeutic method for preventing or treating a vascular indication characterized by a decreased lumen diameter comprising administering to a mammal at risk of or afflicted with said vascular indication, a cytostatic dose of a therapeutic agent, wherein the therapeutic agent is a compound of formula (I):

$$(R^1)(R^2)N(CH_2)_2O$$
 $(Z)$ 
 $R^3$ 
 $(I)$ 

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wherein Z is C=O or a covalent bond; Y is H or  $O(C_1-C_4)$ alkyl,  $R^1$  and  $R^2$  are individually  $(C_1-C_4)$ alkyl or together with N are a saturated heterocyclic group,  $R^3$  is ethyl or chloroethyl,  $R^4$  is H or together with  $R^3$  is  $-CH_2-CH_2$ - or -S-,  $R^5$  is I, OH,  $O(C_1-C_4)$ alkyl or H and  $R^6$  is I,  $O(C_1-C_4)$ alkyl or H with the proviso that when  $R^4$ ,  $R^5$  and  $R^6$  are H,  $R^3$  is not ethyl; or a pharmaceutically acceptable salt thereof.

- 208. (New) The method of claim 207 wherein the cytostatic dose is effective to increase the level of TGF-beta so as to decrease lesion formation or development, inhibit lipid accumulation, increase plaque stability, maintain or increase vessel lumen diameter, or any combination thereof.
- 209. (New) The method of claim 207 wherein the compound of formula (I) is idoxifene, 4-iodotamoxifen, 3-iodotamoxifen, toremifene, or a pharmaceutically acceptable salt thereof.
- 210. (New) The method of claim 207 wherein the administration is systemic.
- 211. (New) The method of claim 207 wherein the compound of formula (I) is administered in a sustained release dosage form.

wherein Z is C=O or a covalent bond; Y is H or  $O(C_1-C_4)$ alkyl,  $R^1$  and  $R^2$  are individually  $(C_1-C_4)$ alkyl, or together with N are a saturated heterocyclic group,  $R^3$  is ethyl or chloroethyl,  $R^4$  is H or together with  $R^3$  is  $-CH_2-CH_2$ ,  $R^5$  is I, OH,  $O(C_1-C_4)$ alkyl or H and  $R^6$  is I,  $O(C_1-C_4)$ alkyl or H; or a pharmaceutically acceptable salt thereof; effective to inhibit stenosis or reduce restenosis of a mammalian vessel following placement of the stent in said vessel.

214.

(New) The stent of claim 212 or 213, wherein  $R^3$  is not ethyl when  $R^4$ ,  $R^5$  and  $R^6$  are H.

215. (New) An intravascular stent comprising a cytostatic amount of a compound of formula (I):

$$(R^1)(R^2)N(CH_2)_2O$$
 $(Z)$ 
 $R^3$ 
 $(I)$ 

wherein Z is C=O or a covalent bond; Y is H or  $O(C_1-C_4)$ alkyl,  $R^1$  and  $R^2$  are individually  $(C_1-C_4)$ alkyl or together with N are a saturated heterocyclic group,  $R^3$  is ethyl or chloroethyl,  $R^4$  is H or together with  $R^3$  is  $-CH_2-CH_2$ - or -S-,  $R^5$  is I, OH or  $O(C_1-C_4)$ alkyl and  $R^6$  is I,  $O(C_1-C_4)$ alkyl or H; or a pharmaceutically acceptable salt thereof; effective to inhibit stenosis or reduce restenosis of a mammalian vessel following placement of the stent in

reduce restenosis of a mammalian vessel following placement of the stent in said vessel.

- 216. (New) The intravascular stent of claim 212, 213 or 215 that is adapted to maintain expanded vessel luminal area following angioplasty.
- 217. (New) The intravascular stent of claim 216 wherein the compound of formula (I) is in a sustained release dosage form.
- 218. (New) The intravascular stent of claim 216 wherein the matrix of the stent comprises the compound of formula (I).
- 219. (New) The intravascular stent of claim 212, 213 or 215 wherein the stent comprises a coating comprising the compound of formula (I).
- 220. (New) The intravascular stent of claim 219 wherein the coating is biodegradable.
- 221. (New) The intravascular stent of claim 219 wherein the coating is porous or permeable to the inhibitor.
- 222. (New) The therapeutic stent of claims 212, 213 or 215 wherein the matrix of the stent is formed from a porous or permeable non-biodegradable material.
- 223. (New) The therapeutic stent of claim 212, 213 or 215 in which the intravascular stent comprises metal or plastic.
- 224. (New) The therapeutic stent of claim 212, 213 or 215 wherein the matrix is formed from a biodegradable material.

225. (New) A therapeutic method comprising inhibiting vascular smooth muscle cell proliferation comprising administering to a mammal an effective cytostatic antiproliferative amount of a compound of formula (I):

$$(R^1)(R^2)N(CH_2)_2O$$
 $(Z)$ 
 $R^3$ 
 $(I)$ 

By

wherein Z is C= O or a covalent bond; Y is H or  $O(C_1-C_4)$ alkyl,  $R^1$  and  $R^2$  are individually  $(C_1-C_4)$ alkyl or together with N are a saturated heterocyclic group,  $R^3$  is ethyl or chloroethyl,  $R^4$  is H or together with  $R^3$  is  $-CH_2-CH_2$ - or -S-,  $R^5$  is I, OH,  $O(C_1-C_4)$ alkyl or H and  $R^6$  is I,  $O(C_1-C_4)$ alkyl or H; or a pharmaceutically acceptable salt thereof, wherein the administration is by placement of a vascular shunt or intravascular stent comprising said compound.

- 226. (New) The method of claim 225 wherein the compound is droloxifene, raloxifene, toremefine, tamoxifen, idoxifene, or a pharmaceutically acceptable salt thereof.
- 227. (New) The method of claim 225 wherein the shunt or stent matrix is impregnated with the compound of formula (I).
- 228. (New) The method of claim 227 wherein the shunt or stent comprises a coating incorporating said compound of formula (I).

- 229. (New) The method of claim 225 wherein the shunt or stent comprises a coating incorporating said compound of formula (I).
- 230. (New) The method of claim 227, 228, or 229 wherein said matrix or said coating is biodegradable.

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231. (New) A therapeutic method for treating a condition selected from the group consisting of arteriosclerosis and small vessel disease, comprising administering to a mammal afflicted with said condition, an effective amount of a compound of formula (I):

$$(R^1)(R^2)N(CH_2)_2O$$
 $(Z)$ 
 $R^3$ 
 $(I)$ 

wherein Z is C=O or a covalent bond; Y is H or  $O(C_1-C_4)$ alkyl,  $R^1$  and  $R^2$  are individually  $(C_1-C_4)$ alkyl or together with N are a saturated heterocyclic group,  $R^3$  is ethyl or chloroethyl,  $R^4$  is H,  $R^5$  is I,  $O(C_1-C_4)$ alkyl or H and  $R^6$  is I,  $O(C_1-C_4)$ alkyl or H with the proviso that when  $R^4$ ,  $R^5$ , and  $R^6$  are H,  $R^3$  is not ethyl; or a pharmaceutically acceptable salt thereof.

232. (New) A method of treating diabetic retinopathy by increasing the level of TGF-beta in a mammal in need thereof, comprising administering an effective amount of a compound of formula (I):